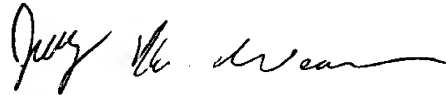


30. (Amended) A method according to claim 1 [or 2] wherein said disease or disorder is chosen from the group consisting of cancer, hyperplasia, restenosis, cardiac hypertrophy, immune disorders and inflammation.

REMARKS

Applicants believe that all pending claims are allowable and respectfully requests a Notice of Allowance for this application from the Examiner. Should the Examiner believe that a telephone conference would expedite the prosecution of this application, the undersigned can be reached at the telephone number set out below.

Respectfully submitted,
BEYER WEAVER & THOMAS, LLP

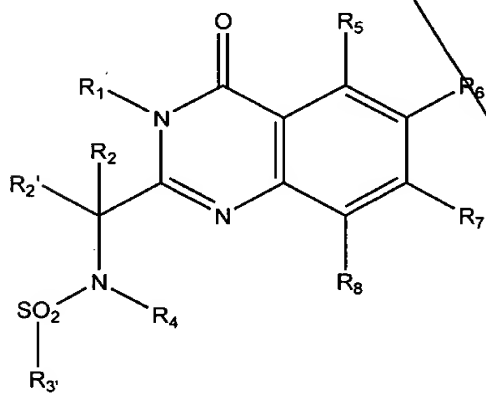
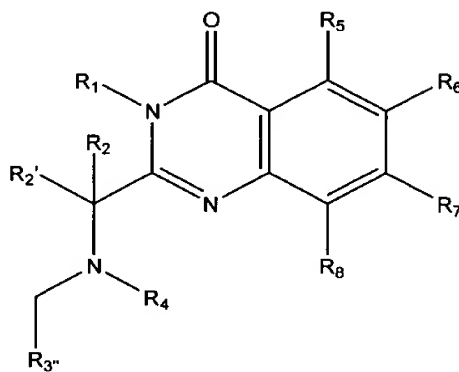
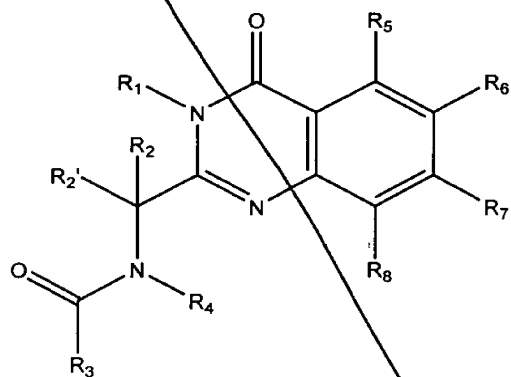


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Reg. No. 31,314

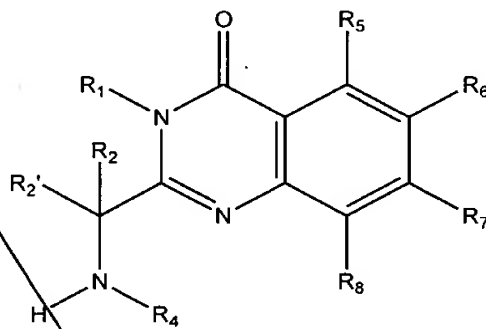
P.O. Box 778
Berkeley, CA 94704-0778

APPENDIX OF PENDING CLAIMS

1. A method of treating cellular proliferative diseases comprising administering a compound chosen from the group consisting of:



and



wherein:

R₁ is chosen from hydrogen, alkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, and substituted alkylheteroaryl;

R₂ and R₂' are independently chosen from hydrogen, alkyl, oxaalkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, and substituted alkylheteroaryl; or R₂ and R₂' taken together form a 3- to 7-membered ring;

R₃ is chosen from hydrogen, alkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, substituted alkylheteroaryl, oxaalkyl, oxaalkylaryl, substituted oxaalkylaryl, R₁₅O- and R₁₅-NH-;

R₃ is chosen from hydrogen, alkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, substituted alkylheteroaryl, and R₁₅-NH-;

R_{3''} is chosen from alkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, and substituted alkylheteroaryl;

R₄ is chosen from hydrogen, alkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, substituted alkylheteroaryl, and R₁₆-alkylene-;

R₅, R₆, R₇ and R₈ are independently chosen from hydrogen, alkyl, alkoxy, halogen, fluoroalkyl, nitro, dialkylamino, alkylsulfonyl, alkylsulfonamido, sulfonamidoalkyl, sulfonamidoaryl, alkylthio, carboxyalkyl, carboxamido, aminocarbonyl, aryl and heteroaryl;

R₁₅ is chosen from alkyl, aryl, alkylaryl, heteroaryl, alkylheteroaryl, substituted alkyl, substituted aryl, substituted alkylaryl, substituted heteroaryl, and substituted alkylheteroaryl;

R₁₆ is chosen from alkoxy, amino, alkylamino, dialkylamino, N-heterocyclyl and substituted N-heterocyclyl.

2-3. Canceled

4. (Amended) A method according to claim 1 wherein

R₁ is chosen from hydrogen, alkyl, aryl, substituted alkyl, substituted aryl, heteroaryl, substituted heteroaryl, alkylaryl, substituted alkylaryl and substituted alkylheteroaryl;

R₂ is chosen from hydrogen, alkyl and substituted alkyl;

R₂' is hydrogen;

R₃ is chosen from alkyl, substituted alkyl, alkylaryl, heteroaryl, aryl, substituted aryl, substituted heteroaryl, substituted oxaalkylaryl R₁₅O- and R₁₅-NH-;

R₄ is chosen from alkyl, aryl, alkylaryl, alkylheteroaryl, substituted alkyl, substituted aryl, and R₁₆-alkylene-;

R₅ is hydrogen;

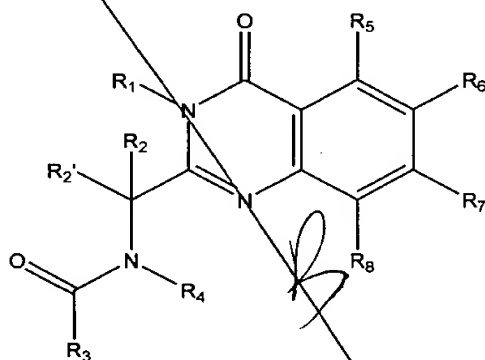
R₆, R₇ and R₈ are independently chosen from hydrogen, halogen, methyl and trifluoromethyl;

R₁₅ is chosen from alkyl, aryl and substituted aryl;

R₁₆ is chosen from alkoxy, amino, alkylamino, dialkylamino and N-heterocyclyl.

5. A method according to claim 4 wherein the stereogenic center to which R₂ and R₂' are attached is of the R configuration.

6. (Amended) A method according to claim 1 comprising administering a compound of formula:



7. A method according to claim 6 wherein R_1 is chosen from hydrogen, lower alkyl, substituted lower alkyl, benzyl, substituted benzyl, phenyl, naphthyl and substituted phenyl.

8. A method according to claim 7 wherein R_1 is chosen from hydrogen, ethyl, propyl, methoxyethyl, naphthyl, phenyl, bromophenyl, chlorophenyl, methoxyphenyl, ethoxyphenyl, tolyl, dimethylphenyl, chlorofluorophenyl, methylchlorophenyl, ethylphenyl, phenethyl, benzyl, chlorobenzyl, methylbenzyl, methoxybenzyl, tetrahydrofuranylmethyl and (ethoxycarbonyl)ethyl.

9. A method according to claim 6 wherein R_2 is chosen from hydrogen, lower alkyl and substituted lower alkyl, and R_2' is hydrogen.

10. A method according to claim 9 wherein R_2 is chosen from hydrogen, methyl, ethyl, propyl, methylthioethyl, aminobutyl, (CBZ)aminobutyl, cyclohexylmethyl, benzyloxymethyl, methylsulfinylethyl, methylsulfinylmethyl, hydroxymethyl, benzyl and indolylmethyl.

11. A method according to claim 6 wherein R_3 is chosen from C_1 - C_{13} alkyl; substituted lower alkyl; phenyl; naphthyl; phenyl substituted with one or more halo, lower alkyl, loweralkoxy, nitro, carboxy, methylenedioxy or trifluoromethyl; biphenyl; benzyl; phenoxymethyl; halophenoxymethyl; phenylvinyl; heteroaryl; heteroaryl substituted with lower alkyl; and benzyloxymethyl.

12. A method according to claim 11 wherein R_3 is chosen from ethyl, propyl, chloropropyl, butoxy, heptyl, butyl, octyl, tridecanyl, (ethoxycarbonyl)ethyl, dimethylaminoethyl, dimethylaminomethyl, phenyl, naphthyl, halophenyl, dihalophenyl, cyanophenyl, halo(trifluoromethyl)phenyl, chlorophenoxymethyl, methoxyphenyl, carboxyphenyl, ethylphenyl, tolyl, biphenyl, methylenedioxyphenyl, methylsulfonylphenyl, methoxychlorophenyl, chloronaphthyl, methylhalophenyl, trifluoromethylphenyl, butylphenyl, pentylphenyl, methylnitrophenyl, phenoxymethyl, dimethoxyphenyl, phenylvinyl, nitrochlorophenyl, nitrophenyl, dinitrophenyl, bis(trifluoromethyl)phenyl, benzyloxymethyl, benzyl, furanyl, benzofuranyl, pyridinyl, indolyl, methylpyridinyl, quinolinyl, picolinyl,

pyrazolyl, and imidazolyl.

13. A method according to claim 6 wherein R_3 is R_{15} -NH- and R_{15} is chosen from lower alkyl; cyclohexyl; phenyl; and phenyl substituted with halo, lower alkyl, loweralkoxy, or lower alkylthio.

14. A method according to claim 13 wherein R_{15} is chosen from isopropyl, butyl, cyclohexyl, phenyl, bromophenyl, dichlorophenyl, methoxyphenyl, ethylphenyl, tolyl, trifluoromethylphenyl and methylthiophenyl.

15. A method according to claim 6 wherein R_4 is chosen from lower alkyl, substituted lower alkyl, cyclohexyl; phenyl substituted with hydroxy, lower alkoxy or lower alkyl; benzyl; heteroarylmethyl; heteroarylethyl; heteroarylpropyl and R_{16} -alkylene-, wherein R_{16} is amino, lower alkylamino, di(lower alkyl)amino, lower alkoxy, or N-heterocyclyl.

16. A method according to claim 15 wherein R_4 is chosen from methyl, ethyl, propyl, butyl, cyclohexyl, carboxyethyl, carboxymethyl, methoxyethyl, hydroxyethyl, hydroxypropyl, dimethylaminoethyl, dimethylaminopropyl, diethylaminoethyl, diethylaminopropyl, aminopropyl, methylaminopropyl, 2,2-dimethyl-3-(dimethylamino)propyl, 1-cyclohexyl-4-(diethylamino)butyl, aminoethyl, aminobutyl, aminopentyl, aminoethyl, aminoethoxyethyl, isopropylaminopropyl, diisopropylaminoethyl, 1-methyl-4-(diethylamino)butyl, (t-Boc)aminopropyl, hydroxyphenyl, benzyl, methoxyphenyl, methylmethoxyphenyl, dimethylphenyl, tolyl, ethylphenyl, (oxopyrrolidinyl)propyl, (methoxycarbonyl)ethyl, benzylpiperidinyl, pyridinylethyl, pyridinylmethyl, morpholinylethyl, morpholinylpropyl, piperidinyl, azetidinylmethyl, azetidinypropyl, pyrrolidinylethyl, pyrrolidinylpropyl, piperidinylmethyl, piperidinylethyl, imidazolylpropyl, imidazolylethyl, (ethylpyrrolidinyl)methyl, (methylpyrrolidinyl)ethyl, (methylpiperidinyl)propyl, (methylpiperazinyl)propyl, furanylmethyl and indolylethyl.

17. A method according to claim 6 wherein R_1 is chosen from lower alkyl, benzyl, substituted benzyl and substituted phenyl; R_2 is chosen from hydrogen, alkyl, substituted lower alkyl and benzyl; R_2' is hydrogen;

R₃ is chosen from substituted phenyl and naphthyl;

R₄ is chosen from substituted alkyl and R₁₆-alkylene-;

R₅ is hydrogen or halo

R₆ is hydrogen, methyl or halo;

R₇ is hydrogen, halo, methyl or trifluoromethyl;

R₈ is hydrogen or halo;

R₁₆ is chosen from di(lower alkylamino), (lower alkyl)amino, amino, N-heterocyclyl and substituted N-heterocyclyl.

Sub
C'
18. (Amended) A method according to claim 1 wherein

R₁ is benzyl or halobenzyl;

R₂ is chosen from ethyl and propyl;

R₂' is hydrogen;

R₃ is substituted phenyl;

R₃' is substituted phenyl;

R₃'' is substituted phenyl;

R₄ is (CH₂)_m OH or (CH₂)_p R₁₆ wherein m is 2 or 3 and p is 1-3;

R₅ is hydrogen;

R₆ is hydrogen;

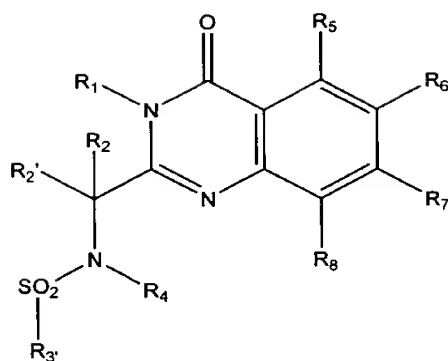
R₇ is halo;

R₈ is hydrogen;

R₁₆ is chosen from amino, propylamino, and azetidiny.

19. A method according to claim 18 wherein the stereogenic center to which R₂ and R₂' are attached is of the R configuration.

20. (Amended) A method according to claim 1 comprising administering a compound of formula:



21. A method according to claim 20 wherein:

R₁ is chosen from hydrogen, lower alkyl, substituted lower alkyl, benzyl, substituted benzyl, phenyl, naphthyl and substituted phenyl;

R₂ is chosen from hydrogen, lower alkyl and substituted lower alkyl and R₂' is hydrogen;

R₃' is chosen from C₁-C₁₃ alkyl, phenyl; naphthyl; phenyl substituted with halo, lower alkyl, lower alkoxy, nitro, methylenedioxy, or trifluoromethyl; biphenyl, benzyl and heteroaryl; and

R₄ is chosen from lower alkyl, substituted lower alkyl, cyclohexyl; phenyl substituted with hydroxy, lower alkoxy or lower alkyl; benzyl; heteroarylmethyl; heteroarylethyl; heteroarylpropyl and R₁₆-alkylene, wherein

R₁₆ is amino, (lower alkyl)amino, di(lower alkyl)amino, lower alkoxy, or N-heterocyclyl.

22. A method according to claim 20 wherein

R₁ is chosen from lower alkyl, benzyl, substituted benzyl and substituted phenyl;

R₂ is hydrogen or lower alkyl;

R₂' is hydrogen;

R₃ is chosen from substituted phenyl and naphthyl;

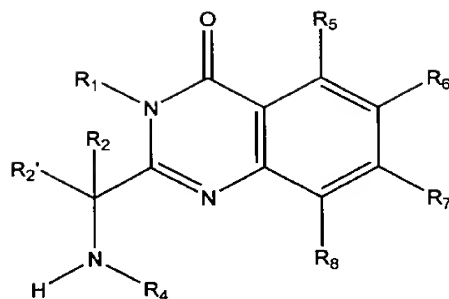
R₄ is R₁₆-alkylene-, hydroxy lower alkyl or carboxy lower alkyl;

R₆ and R₇ are chosen from hydrogen and halo;

R₅ and R₈ are hydrogen;

R₁₆ is chosen from di(lower alkylamino), (lower alkyl)amino, amino, piperidinyl, azetidiny, pyrrolidinyl and morpholinyl.

23. (Amended) A method according to claim 1 comprising administering a compound of formula:



24. A method according to claim 23 wherein:

R₁ is chosen from hydrogen, lower alkyl, substituted lower alkyl, benzyl, substituted benzyl, phenyl, naphthyl and substituted phenyl;

R₂ is chosen from hydrogen, lower alkyl and substituted lower alkyl and R₂' is hydrogen; and

R₄ is chosen from lower alkyl, cyclohexyl; phenyl substituted with hydroxy, lower alkoxy or lower alkyl; benzyl; heteroarylmethyl; heteroarylethyl; heteroarylpropyl and R₁₆-alkylene, wherein R₁₆ is di(lower alkyl)amino, alkylamino, amino, lower alkoxy, or N-heterocyclyl.

25. A method according to claim 23 wherein

R₁ is chosen from lower alkyl, benzyl, substituted benzyl and substituted phenyl;

R₂ is hydrogen or lower alkyl;

R₂' is hydrogen;

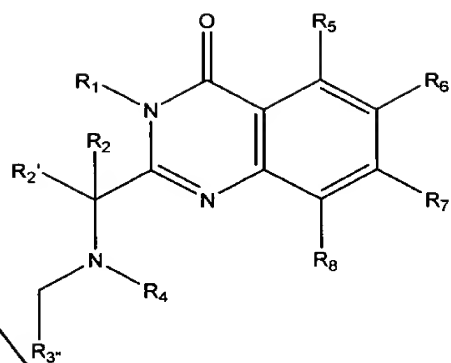
R₄ is R₁₆-alkylene-;

R₆ and R₇ are chosen from hydrogen and halo;

R₅ and R₈ are hydrogen;

R₁₆ is chosen from di(lower alkylamino), (lower alkyl)amino, amino, pyrrolidinyl, piperidinyl, imidazolyl and morpholinyl.

26. (Amended) A method according to claim 1 comprising administering a compound of formula:



27. A method according to claim 26 wherein:

- R₁ is chosen from hydrogen, lower alkyl, substituted lower alkyl, benzyl, substituted benzyl, phenyl, naphthyl and substituted phenyl;
- R₂ is chosen from hydrogen, lower alkyl and substituted lower alkyl and R₂' is hydrogen;
- R₃'' is chosen from C₁-C₁₃ alkyl; substituted lower alkyl; phenyl; naphthyl; phenyl substituted with halo, lower alkyl, lower alkoxy, nitro, methylenedioxy, or trifluoromethyl; biphenyl; benzyl and heterocyclyl; and
- R₄ is chosen from lower alkyl, substituted lower alkyl; cyclohexyl; phenyl substituted with hydroxy, lower alkoxy or lower alkyl; benzyl; substituted benzyl; heterocyclyl; heteroarylmethyl; heteroarylethyl; heteroarylpropyl and R₁₆-alkylene, wherein R₁₆ is di(lower alkyl)amino, (lower alkyl)amino, amino, lower alkoxy, or N-heterocyclyl.

28. A method according to claim 27 wherein

R₁ is chosen from lower alkyl, benzyl, substituted benzyl and substituted phenyl;

R₂ is hydrogen or lower alkyl;

R₂' is hydrogen;

R₃ is chosen from substituted phenyl, heterocyclyl and naphthyl;

R₄ is chosen from substituted benzyl, heterocyclyl substituted lower alkyl and R₁₆-alkylene-;

R₆ and R₇ are chosen from hydrogen and halo;

R₅ and R₈ are hydrogen;

R₁₆ is chosen from di(lower alkylamino), (lower alkyl)amino, amino, pyrrolidinyl, azetidiny, piperidinyl, imidazolyl and morpholinyl.

29. A method according to claim 28 wherein

R₁ is benzyl;

R₂ is ethyl;

R₂' is hydrogen;

R₃ is chosen from halophenyl, polyhalophenyl, tolyl, dimethylphenyl, methoxyphenyl, dimethoxyphenyl, cyanophenyl, trifluoromethylphenyl, trifluoromethoxyphenyl, bis(trifluoromethyl)phenyl, carboxyphenyl, t-butylphenyl, methoxycarbonylphenyl, piperidinyl and naphthyl;

R₄ is chosen from substituted benzyl, piperidinyl, hydroxy (lower alkyl) and R₁₆-alkylene-;

R₆ and R₇ are chosen from hydrogen and halo;

R₅ and R₈ are hydrogen;

R₁₆ is chosen from dimethylamino, amino, pyrrolidinyl and piperidinyl.

30. (Amended)

A method according to claim 1 wherein said disease or disorder is chosen from the group consisting of cancer, hyperplasia, restenosis, cardiac hypertrophy, immune disorders and inflammation.

31-59. Canceled